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  - L4 ANSWER 1 OF 8 HCAPLUS COPYRIGHT 2000 ACS
  - AN 1999:396638 HCAPLUS
  - DN 131:212774
  - TI Mapping of the interleukin-10/interleukin-10 receptor combining site using structurally different peptide scans
  - AU Reineke, Ulrich; Sabat, Robert; Volk, Hans-Dieter; Schneider-Mergener, Jens
  - CS Institut fur Medizinische Immunologie, Universitatsklinikum Charite, Humboldt-Universitat zu Berlin, Berlin, 10098, Germany
  - SO Pept. Proc. Am. Pept. Symp., 15th (1999), Meeting Date 1997, 533-534. Editor(s): Tam, James P.; Kaumaya, Pravin T. P. Publisher: Kluwer, Dordrecht, Neth. CODEN: 67UCAR
  - DT Conference
  - LA English
  - AB The authors describe strategies for the mapping of putative interleukin-10/interleukin-10 receptor combining site: (1)the detection of low affinity protein-peptide interactions and (2) the use of overlapping peptide scans with peptides of different length.
  - L4 ANSWER 2 OF 8 HCAPLUS COPYRIGHT 2000 ACS

AN 1998:250487 HCAPLUS

DN 129:26823

TI Mapping of the interleukin-10/interleukin-10 receptor combining site

AU Reineke, Ulrich; Sabat, Robert; Volk, Hans-Dieter; Schneider-Mergener,

CS Institut fur Medizinische Immunologie, Universitatsklinikum Charite, Humboldt-Universitat zu Berlin, Berlin, 10098, Germany

SO Protein Sci. (1998), 7(4), 951-960 CODEN: PRCIEI; ISSN: 0961-8368

PB Cambridge University Press

DT Journal

LA English

AΒ The discontinuous interleukin-10 (IL-10)/interleukin-10 receptor (IL-10R) combining site was mapped using sets of overlapping peptides derived from both binding partners bound to continuous cellulose membranes. Low affinity binding of single regions of the discontinuous contact sites on IL-10 and IL-10R could be identified due to (1) high peptide d. on the membrane support, (2) incubation with high protein concns., (3) indirect immunodetection of the ligates after electrotransfer onto polyvinylene difluoride membranes, and (4) use of highly overlapping peptide scans of different length (6-mers and 15-mers). The single binding regions identified for each protein species are sepd. in the protein sequences, but form continuous areas on the surface of IL-10 (x-ray structure) and IL-10R (computer model). Furthermore, 4 epitopes of neutralizing anti-IL-10 and anti-IL-10R antibodies were mapped and overlap with these binding regions. Sol. peptides (15-19-mers) each spanning one of the 3 identified IL-10-derived receptor binding regions displayed no affinity to IL-10R as expected, whereas a peptide (35-mer) comprising two of these regions had considerably higher binding activity. The data are consistent with a previously published computer model of the IL-10/IL-10R complex. This approach should be generally applicable for the mapping of non-linear protein-protein contact sites.

L4 ANSWER 3 OF 8 HCAPLUS COPYRIGHT 2000 ACS

AN 1998:151218 HCAPLUS

DN 128:150400

TI Complete genome sequence of the methanogenic archaeon, Methanococcus jannaschii

IN Bult, Carol J.; White, Owen R.; Smith, Hamilton O.; Woese, Carl R.;
Venter, J. Craig

PA Institute for Genomic Research, USA; Board of Trustees of the University of Illinois; Johns Hopkins University School of Medicine

SO PCT Int. Appl., 615 pp. CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

W: CA, JP

RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE PRAI US 1996-24428 19960822

AB The present application describes the complete 1.66-megabase pair genome sequence of an autotrophic archaeon, Methanococcus jannaschii, and its 58-and 16-kilobase pair extrachromosomal elements. Also described are 1738

predicted protein-coding genes. Computer-readable media for the storage, search and retrieval of the M. jannaschii genome sequence are also provided.

- ANSWER 4 OF 8 HCAPLUS COPYRIGHT 2000 ACS L4
- 1998:26596 HCAPLUS AN
- 128:126882 DN
- Identification of functional domains on human interleukin 10 ΤI
- Gesser, Borbala; Leffers, Henrik; Jinquan, Tan; Vestergaard, Christian; ΑU Kirstein, Nicka; Sindet-Pedersen, Steen; Lindkaer Jensen, Steen; Thestrup-Pedersen, Kristian; Gronhoj Larsen, Christian
- Department Dermatology, Marselisborg Hospital, University Aarhus, Aarhus, CS DK-8000, Den.
- Proc. Natl. Acad. Sci. U. S. A. (1997), 94(26), 14620-14625 SO CODEN: PNASA6; ISSN: 0027-8424
- National Academy of Sciences PB
- DTJournal
- English LA
- Interleukin 10 (IL-10) is a recently described natural endogenous AB immunosuppressive cytokine that has been identified in humans, mice, and other organisms. Human IL-10 (hIL-10) has high homol. with murine IL-10 (mIL-10) as well as with an Epstein-Barr virus genome product BCRFI. viral IL-10 (vIL-10) shares a no. of activities with hIL-10. IL-10 affects chemokine biol., because human IL-10 inhibits chemokine prodn. and is a specific chemotactic factor for CD8+ T cells. It suppresses the ability of CD4+ T cells, but not CD8+ T cells, to migrate in response to IL-8. A nonapeptide (IT9302) with complete homol. to a sequence of hIL-10 located in the C-terminal portion (residues 152-160) of the cytokine was found to possess activities that mimic some of those of hIL-10. These are: (1) inhibition of IL-1.beta.-induced IL-8 prodn. by peripheral blood mononuclear cell, (2) inhibition of spontaneous IL-8 prodn. by cultured human monocytes, (3) induction of IL-1 receptor antagonist protein prodn. by human monocytes, (4) induction of chemotactic migration of CD8+ human T lymphocytes in vitro, (5) desensitization of human CD8+ T cells resulting in an unresponsiveness toward rhIL-10-induced chemotaxis, (6) suppression of the chemotactic response of CD4+ T human lymphocytes toward IL-8, (7) induction of IL-4 prodn. by cultured normal human CD4+ T cells, (8) down-regulation of tumor necrosis factor-.alpha. prodn. by CD8+ T cells, and (9) inhibition of class II major histocompatibility complex antigen expression on IFN-.gamma.-stimulated human monocytes. Another nonapeptide (IT9403) close to the N-terminal part of hIL-10 did not reveal cytokine synthesis inhibitory properties, but proved to be a regulator of mast cell proliferation. Thus, the authors identified 2 functional domains of IL-10 exerting different IL-10 like activities, an observation that suggests that relatively small segments of these signal proteins are responsible for particular biol. functions.
- ANSWER 5 OF 8 HCAPLUS COPYRIGHT 2000 ACS L4
- 1997:499202 HCAPLUS AN
- 127:160576 DN
- TI Synthetic IL-10 analogs
- Gronhoj Larsen, Christian; Gesser, Borbala IN
- Steeno Research Group A/S, Den.; Gronhoj Larsen, Christian; Gesser, PA Borbala
- PCT Int. Appl., 101 pp. so

CODEN: PIXXD2

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Patent
DT
LA
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FAN.CNT 1
                                                            DATE
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                                                            19960118
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                       A1
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PRAI DK 1996-9629
                       19960118
     WO 1996-DK29
                       19970116
     WO 1997-DK21
     The invention relates to use of a substance or polypeptide according to
ΑB
     the formula: X1-X2-X3-Thr-X4-Lys-X5-Arg-X6, wherein X1 is Ala or Gly, X2
     is Tyr or Phe, X3, X4 and X5 are independently selected from the group
     consisting of Met, Ile, Leu and Val; and X6 is selected from the group
     consisting of Asp, Gln and Glu, optionally at least one of X1, X2, X3, X4,
     X5 and X6 is independently substituted with non-natural or unusual amino
      acids and/or the peptide is cyclized and/or the peptide is stabilized
      and/or the amino terminal amino acid residue is acylated and/or the
      carboxy terminal amino acid residue is amidated, and peptidomimetics
      modelled on the basis of the above formula for the prepn. of a
      pharmaceutical compn. for the redn. of TNF.alpha. prodn. These peptides
      inhibit IL-8 prodn. by human monocytes and IL-1.beta.-induced IL-8 prodn.
      in peripheral blood mononuclear cells, chemotactic response of CD4+ T
      lymphocytes, chemotactic response of human monocytes towards MCAF/MCP-1,
      class II MHC mol. expression on human monocytes, TNF-.alpha. prodn. in
      mixed leukocyte, etc. The IL-10 analogs also induce prodn. of IL-1
      receptor antagonist protein in human monocytes, chemotaxis of CD8+ T
      lymphocytes, IL-4 prodn. in CD4+ T lymphocytes, and modulate LPS-induced
      shock and leukopenia and bile acid-induced pancreatitis. These peptides
      are useful for treating conditions related to disturbance of a cytokine
      system, or diseases where macrophages/T-lymphocyte-mediated immune
      reactions are considered pathogenetically important.
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ANSWER 6 OF 8 HCAPLUS COPYRIGHT 2000 ACS
L4
AN
     1996:516805 HCAPLUS
DN
     125:266912
     Complete genome sequence of the methanogenic archaeon, Methanococcus
TI
     jannaschii
ΑU
     Bult, Carol J.; White, Owen; Olsen, Gary J.; Zhou, Lixin; Fleischmann,
     Robert D.; Sutton, Granger G.; Blake, Judith A.; FitzGerald, Lisa M.;
     Clayton, Rebecca A.; et al.
     TIGR, Rockville, MD, 20850, USA
CS
     Science (Washington, D. C.) (1996), 273(5278), 1058-1073
     CODEN: SCIEAS; ISSN: 0036-8075
DT
     Journal
LA
     English
AB
     The complete 1.66-megabase pair genome sequence of an autotrophic
     archaeon, Methanococcus jannaschii, and its 58- and 16-kilobase pair
     extrachromosomal elements were detd. by whole-genome random sequencing.
     total of 1738 predicted protein-coding genes were identified; however,
     only a minority of these (38%) could be assigned a putative cellular role
     with high confidence. Although the majority of genes related to energy
     prodn., cell division, and metab. in M. jannaschii are most similar to
     those found in bacteria, most of the genes involved in transcription,
     translation, and replication in M. jannaschii are more similar to those
     found in eukaryotes.
     ANSWER 7 OF 8 HCAPLUS COPYRIGHT 2000 ACS
L4
ΑN
     1996:133072 HCAPLUS
DN
     124:173458
ΤI
     Interleukin 10 agonist polypeptides as immunomodulators
     Groenhoej Larsen, Christian; Gesser, Borbala
IN
PA
     Nycomed Dak A/s, Den.
so
     PCT Int. Appl., 98 pp.
     CODEN: PIXXD2
DT
     Patent
     English
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Searched by David Schreiber 308-4292

NO 1997-20

19970103

19970305

Α

NO 9700020

PRAI DK 1994-800 19940705 WO 1995-DK227 19950607

OS MARPAT 124:173458

AB This invention relates to a polypeptide other than human interleukin 10 which has at least one of the following properties: a) induces inhibition of spontaneous IL-8 prodn. by human monocytes, b) induces inhibition of IL-1.beta. induced IL-8 prodn. by human peripheral blood mononuclear cells (PBMC), c) induces prodn. of interleukin-1 receptor antagonistic protein (IRAP) by human monocytes, d) induces chemotactic migration of CD8+ human T lymphocytes in vitro, e) desensitizes human CD8+ T cells resulting in an ... unresponsiveness towards rhIL-10, f) suppresses the chemotactic response of CD4+ T human lymphocytes towards IL-8, g) suppresses the chemotactic response of human monocytes towards MCAF/MCP-1, h) does not inhibit class II MHC mol. expression on human monocytes, in contrast to human IL-10, i) induces the prodn. of IL-4 by cultured normal human CD4+ T cells, j) rescued the TNF.alpha. prodn. in human mixed leukocyte reaction. particular, the invention relates to the nonapeptide Ala-Tyr-Met-Thr-Met-Lys-Ile-Arg-Asn and analogs and variants thereof. In example, the functions of synthetic polypeptide IT9302 were characterized, and antibody against the synthetic polypeptide IT9302 was prepd.

L4 ANSWER 8 OF 8 HCAPLUS COPYRIGHT 2000 ACS

AN 1994:453312 HCAPLUS

DN 121:53312

TI Secondary structures of lipid-associating peptides: a Fourier transform infrared study

AU Zhong, Qi; Clark-Lewis, Ian; Cushley, Robert J.

CS Simon Fraser Univ., Burnaby, BC, Can.

SO Pept. Res. (1994), 7(2), 99-106 CODEN: PEREEO; ISSN: 1040-5704

DT Journal

LA English

AB Four peptides from 20 to 28 residues in length were studied by Fourier transform IR (FTIR) spectroscopy in soln. and in complexes with dimyristoylphosphatidylcholine (DMPC). The four peptides included the 20-residue lipid-assocg. peptide, LAP-20, which was predicted to form an amphipathic helical structure in the presence of lipids, and three other peptides whose sequences had less amphipathic helix-forming properties. The complexes were shown by electron microscopy to be discoidal in shape with mean diams. of 21-27 nm. At the concns. used for IR, the peptides appeared to form oligomers consisting of intermol. .beta.-sheets. In the presence of lipids, the amt. of .beta.-structure decreased; however, amts. of .beta.-structure were still approx. equal to amts. of .alpha.-helix. The IR results for LAP-20 contradicted previous CD results that predicted 50%-90% .alpha.-helix in DMPC complexes. Convex constraint anal. (CCA) deconvolution of the CD (CD) spectrum to est. secondary structures predicted amts. of helix similar to those predicted by IR, but there was still substantial disagreement between IR and CD ests. of other secondary structures. For LAP-20 in complexes, CD predicted random structure. Possible physiol. consequences of partial disordering of peptide structures are discussed.

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